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99) US

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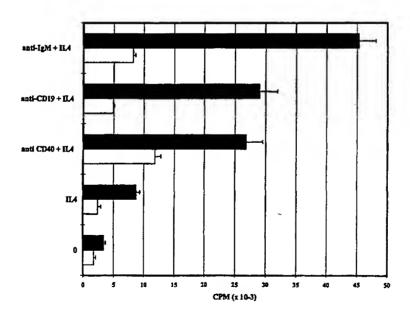
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#### (57) Abstract

Soluble, secreted tumor necrosis factor receptor polypeptides, polynucleotides encoding the polypeptides, and related compositions and methods are disclosed. The polypeptides comprise one cysteine-rich repeat that is homologous to other tumor necrosis factor receptors, such as transmembrane activator and CAML-interactor (TACI). The polypeptides may be used for detecting ligands, agonists and antagonists. The polypeptides may also be used in methods that modulate B cell activation.

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## INTERNATIONAL SEARCH REPORT

Int Alional Application No PCT/US 00/00396

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A CLASS	FIGATION OF SUBJECT MATTER C12N15/11 'C07K14/705 A61K3	95	
	o International Patent Classification (IPC) or to both national clas	ssification and IPC	
	SEARCHED ocumentation searched (classification system followed by classification system followed by classifi	lication symbole:	
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	lata base consulted during the international search (name of dat	a base and. where practical, se	earch (erms used)
CHEM A	BS Data, BIOSIS		
	ENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the	e relevant passages	Relevant to claim No.
х	WO 98 39361 A (ST. JUDE CHILDRE	EN'S	56-63
	RESEARCH HOSPITAL) 11 September 1998 (1998-09-11)		
	cited in the application		
	the whole document		
X	G-U VON BÜLOW AND R J BRAM: "N	NF-AT	56-63
	activation induced by a CAML-in member of the tumor necrosis fa	nteracting	
	receptor superfamily"	ictor	
	SCIENCE., vol. 278, 3 October 1997 (1997-	-1002)	
i	pages 138-141, XP002140938	-10-03),	
	AAAS. LANCASTER, PA., US cited in the application		
İ	the whole document		
		,	
		-/	
X Furth	er documents are listed in the continuation of box C.	X Patent family mer	mbers are listed in annex.
	egories of cited documents :	"T" later document publish	ed after the international filing date
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#### INTERNATIONAL SEARCH REPORT

In. ational Application No PCT/US 00/00396

		PCT/US 00/00396
C.(Continu	uation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Jategory -	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 98 18921 A (HUMAN GENOME SCIENCES) 7 May 1998 (1998-05-07) cited in the application the whole document	1-63
A	EP 0 869 180 A (SMITH KLINE BEECHAM CORPORATION) / October 1998 (1998-10-07) the whole document	1-63
Ţ	the whole document  J A GROSS ET AL.: "TACI and BCMA are receptors for a TNF homologue implicated in B-cell autoimmune disease" NATURE., vol. 404, 27 April 2000 (2000-04-27), pages 995-999, XP002140939 MACMILLAN JOURNALS LTD. LONDON., GB ISSN: 0028-0836 the whole document	1-63

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## INTERNATIONAL SEARCH REPORT

Information on patent family members

In. atlanal Application No PCT/US 00/00396

Patent document cited in search report	11	Publication date		atent family nember(s)		Publication date
WO 9839361	А	11-09-1998	US AU EP	5969102 6685498 0964874	Α	19-10-1999 22-09-1998 22-12-1999
WO 9818921	A	07-05-1998	AU BR EP	7674596 9612752 0939804	A	22-05-1998 18-01-2000 08-09-1999
EP 869180	A	07-10-1998	CA JP JP 20	2232743 10323194 000060580	A	02-10-1998 08-12-1998 29-02-2000